

Welcome to STN International! Enter x:x

LOGINID:sssptal611sxp

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

* * * * * Welcome to STN International * * * * *

NEWS 1		Web Page URLs for STN Seminar Schedule - N. America
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NEWS 4	Feb 24	TEMA now available on STN
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NEWS 18	May 15	Supporter information for ENCOMPAT and ENCOMPLIT updated
NEWS 19	May 19	Simultaneous left and right truncation added to WSCA
NEWS 20	May 19	RAPRA enhanced with new search field, simultaneous left and right truncation
NEWS 21	Jun 06	Simultaneous left and right truncation added to CBNB
NEWS 22	Jun 06	PASCAL enhanced with additional data
NEWS 23	Jun 20	2003 edition of the FSTA Thesaurus is now available
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NEWS HOURS		STN Operating Hours Plus Help Desk Availability
NEWS INTER		General Internet Information
NEWS LOGIN		Welcome Banner and News Items
NEWS PHONE		Direct Dial and Telecommunication Network Access to STN
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Enter NEWS followed by the item number or name to see news on that specific topic.

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* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 09:42:41 ON 28 JUL 2003

=> file reg

COST IN U.S. DOLLARS

SINCE FILE ENTRY TOTAL
SESSION

FULL ESTIMATED COST

0.21 0.21

FILE 'REGISTRY' ENTERED AT 09:42:46 ON 28 JUL 2003

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STRUCTURE FILE UPDATES: 25 JUL 2003 HIGHEST RN 555152-78-8

DICTIONARY FILE UPDATES: 25 JUL 2003 HIGHEST RN 555152-78-8

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:

<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=>

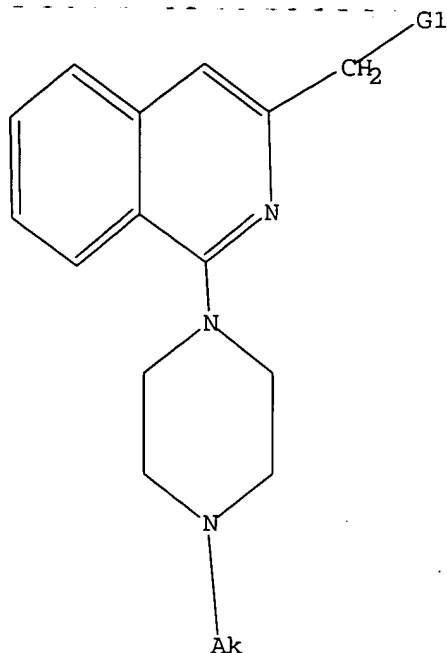
Uploading 09852850.1

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR



G1 Cb,Cy,Hy

Structure attributes must be viewed using STN Express query preparation.

=> s l1 sss full

FULL SEARCH INITIATED 09:43:17 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 1033 TO ITERATE

100.0% PROCESSED 1033 ITERATIONS

10 ANSWERS

SEARCH TIME: 00.00.01

L2 10 SEA SSS FUL L1

=> file caold

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

148.15

148.36

FILE 'CAOLD' ENTERED AT 09:43:23 ON 28 JUL 2003

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FILE COVERS 1907-1966

FILE LAST UPDATED: 01 May 1997 (19970501/UP)

This file contains CAS Registry Numbers for easy and accurate substance identification. Title keywords, authors, patent assignees, and patent information, e.g., patent numbers, are now searchable from 1907-1966. TIFF images of CA abstracts printed between 1907-1966 are available in the PAGE display formats.

This file supports REGISTRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.

=> s l1

REGISTRY INITIATED

Substance data SEARCH and crossover from CAS REGISTRY in progress...
Use DISPLAY HITSTR (or FHITSTR) to directly view retrieved structures.

SAMPLE SEARCH INITIATED 09:43:28 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 66 TO ITERATE

100.0% PROCESSED 66 ITERATIONS 0 ANSWERS
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 833 TO 1807
PROJECTED ANSWERS: 0 TO 0

L3 0 SEA SSS SAM L1

L4 0 L3

=> s l2

L5 0 L2

=> file caold

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.80	149.96

FILE 'CAOLD' ENTERED AT 09:44:30 ON 28 JUL 2003
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FILE COVERS 1907-1966

FILE LAST UPDATED: 01 May 1997 (19970501/UP)

This file contains CAS Registry Numbers for easy and accurate substance identification. Title keywords, authors, patent assignees, and patent information, e.g., patent numbers, are now searchable from 1907-1966. TIFF images of CA abstracts printed between 1907-1966 are available in the PAGE display formats.

This file supports REGISTRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.

=> s l1 sss full

REGISTRY INITIATED

Substance data SEARCH and crossover from CAS REGISTRY in progress...
Use DISPLAY HITSTR (or FHITSTR) to directly view retrieved structures.

FULL SEARCH INITIATED 09:44:50 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 1033 TO ITERATE

100.0% PROCESSED 1033 ITERATIONS
SEARCH TIME: 00.00.01

10 ANSWERS

L6 10 SEA SSS FUL L1

L7 0 L6

=> logy

LOGY IS NOT A RECOGNIZED COMMAND

The previous command name entered was not recognized by the system.
For a list of commands available to you in the current file, enter
"HELP COMMANDS" at an arrow prompt (=>).

=> log y

COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
0.40	298.91

FULL ESTIMATED COST

STN INTERNATIONAL LOGOFF AT 09:45:06 ON 28 JUL 2003

Welcome to STN International! Enter x:x

LOGINID:sssptal611sxp

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

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* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 09:38:26 ON 28 JUL 2003

=> file reg

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

0.21

0.21

FILE 'REGISTRY' ENTERED AT 09:38:37 ON 28 JUL 2003

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STRUCTURE FILE UPDATES: 25 JUL 2003 HIGHEST RN 555152-78-8

DICTIONARY FILE UPDATES: 25 JUL 2003 HIGHEST RN 555152-78-8

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:

<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

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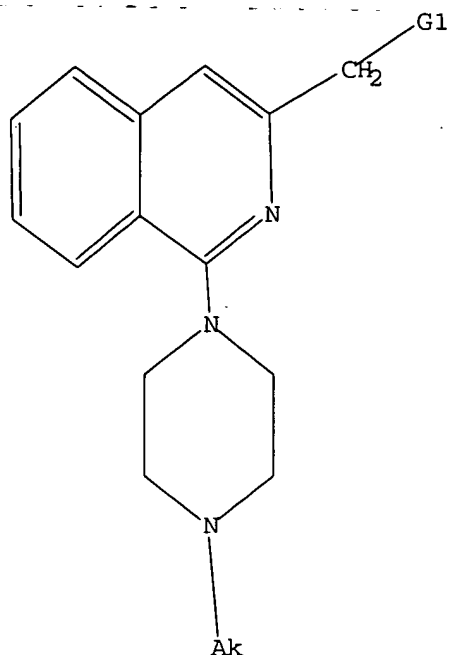
Uploading 09852850.1

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR



G1 Cb,Cy,Hy

Structure attributes must be viewed using STN Express query preparation.

=> s l1

SAMPLE SEARCH INITIATED 09:38:59 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 66 TO ITERATE

100.0% PROCESSED 66 ITERATIONS
SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 833 TO 1807
PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> s l1 sss full

FULL SEARCH INITIATED 09:39:06 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 1033 TO ITERATE

100.0% PROCESSED 1033 ITERATIONS
SEARCH TIME: 00.00.01

10 ANSWERS

L3 10 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
148.15	148.36

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 09:39:15 ON 28 JUL 2003
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FILE COVERS 1907 - 28 Jul 2003 VOL 139 ISS 5
FILE LAST UPDATED: 27 Jul 2003 (20030727/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s l3

L4 4 L3

=> d l4 fbib hitstr abs total

L4 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN
AN 1999:244638 CAPLUS
DN 130:311813
TI Preparation of piperazinyloquinolines and analogs as serotonin antagonists
IN Ueno, Kohshi; Sasaki, Atsushi; Kawano, Koki; Okabe, Tadashi; Kitazawa, Noritaka; Takahashi, Keiko; Yamamoto, Noboru; Suzuki, Yuichi; Matsunaga, Manabu; Kubota, Atsuhiko
PA Eisai Co., Ltd., Japan
SO PCT Int. Appl., 740 pp.
CODEN: PIXXD2
DT Patent
LA Japanese
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9918077	A1	19990415	WO 1998-JP4465	19981002
	W: US				
	RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	JP 2000053647	A2	20000222	JP 1997-284290 A	19971002
				JP 1998-281752	19981002
				JP 1997-284290 A	19971002
				JP 1998-153416 A	19980602
EP 1020445	A1	20000719		EP 1998-945593	19981002
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
				JP 1997-284290 A	19971002
				WO 1998-JP4465 W	19981002
US 6340759	B1	20020122		US 2000-509778	20000331

US 2002013460 A1 20020131

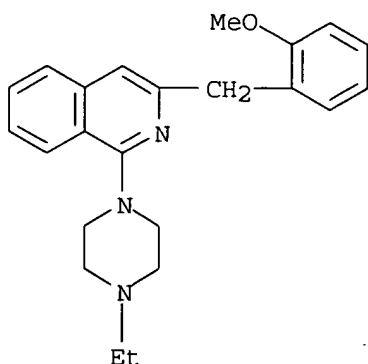
JP 1997-284290 A 19971002
WO 1998-JP4465 W 19981002
US 2001-852850 20010511
JP 1997-284290 A 19971002
WO 1998-JP4465 W 19981002
US 2000-509778 A320000331

OS MARPAT 130:311813

IT 223542-46-9P 223542-47-0P 223551-31-3P
223551-33-5P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(prepn. of piperazinyliisoquinolines and analogs as serotonin antagonists)

RN 223542-46-9 CAPLUS

CN Isoquinoline, 1-(4-ethyl-1-piperazinyl)-3-[(2-methoxyphenyl)methyl]- (9CI)
(CA INDEX NAME)

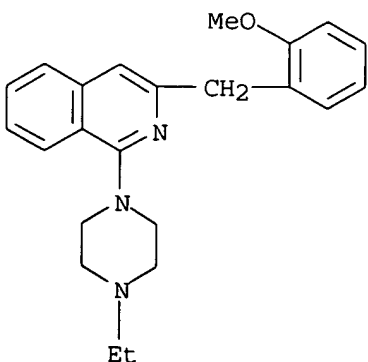
RN 223542-47-0 CAPLUS

CN Isoquinoline, 1-(4-ethyl-1-piperazinyl)-3-[(2-methoxyphenyl)methyl]-, ethanedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 223542-46-9

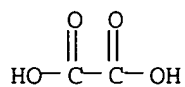
CMF C23 H27 N3 O



CM 2

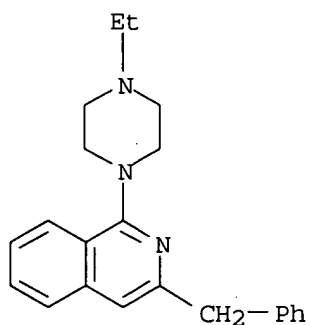
CRN 144-62-7

CMF C2 H2 O4



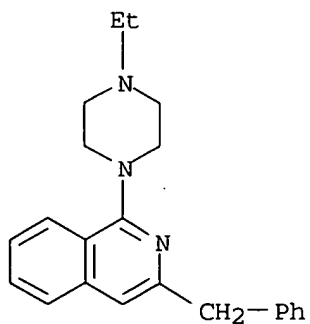
RN 223551-31-3 CAPLUS

CN Isoquinoline, 1-(4-ethyl-1-piperazinyl)-3-(phenylmethyl)- (9CI) (CA INDEX NAME)



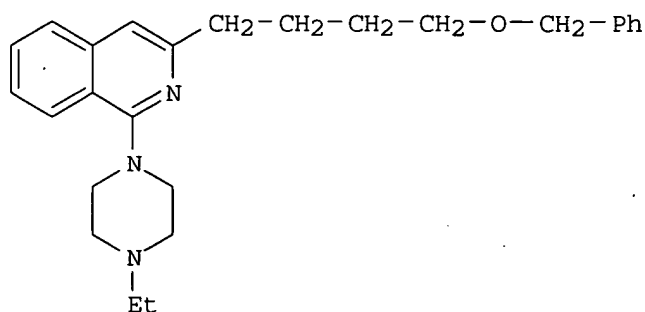
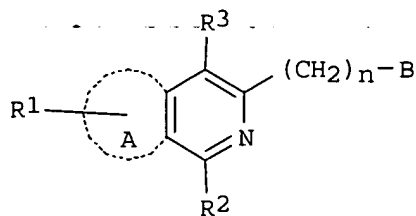
RN 223551-33-5 CAPLUS

CN Isoquinoline, 1-(4-ethyl-1-piperazinyl)-3-(phenylmethyl)-, dihydrochloride (9CI) (CA INDEX NAME)



● 2 HCl

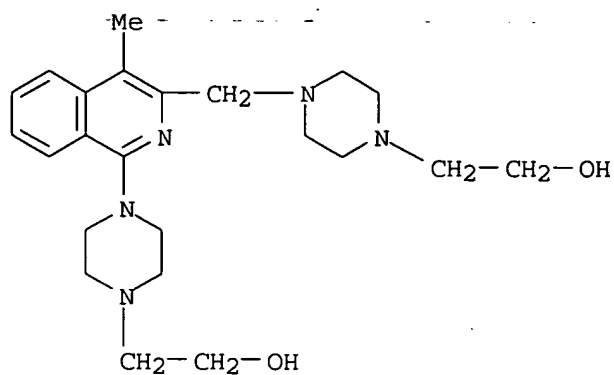
GI



AB The title compds. I [ring A = benzene, pyridine, thiophene or furan ring; B = (un)substituted aryl, etc.; R1 = H, halo, etc.; R2 = 4-morpholinyl, etc.; R3 = H, halo, etc.; n = 0, or 1 - 6] are prepd. I are central muscle relaxing drugs for treating, ameliorating or preventing spastic paralysis or ameliorating myotonia. In an in vitro test for 5HT1 receptor antagonism, the title compd. II showed the Ki value of 21.2 nM.

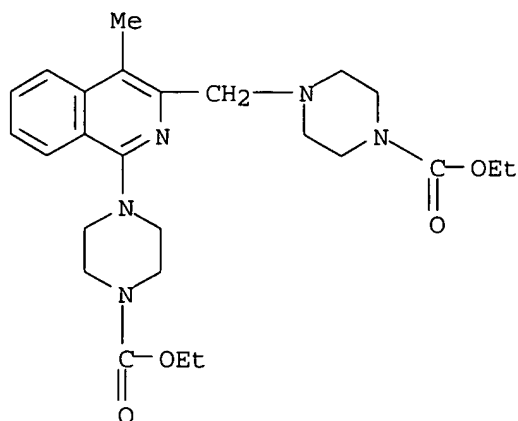
RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN
AN 1972:564414 CAPLUS
DN 77:164414
TI Reactions of 1-chloro-3-chloromethyl-4-methylisoquinoline
AU Nair, M. D.
CS Ciba Res. Cent., Bombay, India
SO Indian Journal of Chemistry (1972), 10(4), 337-40
CODEN: IJOCAP; ISSN: 0019-5103
DT Journal
LA English
IT 14576-16-0P 14576-17-1P 14577-67-4P
RL: SPN (Synthetic preparation); PREP (Preparation)
(prepn. of)
RN 14576-16-0 CAPLUS
CN 1-Piperazineethanol, 4-[[1-[4-(2-hydroxyethyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]- (9CI) (CA INDEX NAME)



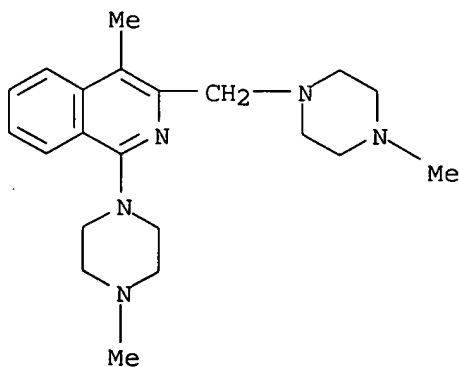
RN 14576-17-1 CAPLUS

CN 1-Piperazinecarboxylic acid, 4-[[1-[4-(ethoxycarbonyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]-, ethyl ester (9CI) (CA INDEX NAME)



RN 14577-67-4 CAPLUS

CN Isoquinoline, 4-methyl-1-(4-methyl-1-piperazinyl)-3-[(4-methyl-1-piperazinyl)methyl]- (8CI, 9CI) (CA INDEX NAME)



GI For diagram(s), see printed CA Issue.

AB With secondary bases 1-chloro-3-(chloromethyl)-4-methylisoquinoline (I) gave mono or disubstitution products in which the Cl in positions 1 or 3, or both was replaced. In 1-chloro-3-[(2-methylpiperidino)-methyl]-4-methylisoquinoline there was NMR evidence for non-equivalence of benzylic methylene protons from the asymmetry of the 2-Me substituent on piperidine. Reaction of I with piperazine gave a bis condensation product, II, with NH₃ and 4-(.gamma.-aminopropyl)morpholine III and IV were obtained, resp. Nitration of I gave the corresponding 5-NO₂ deriv., reaction of which with bases gave mono or disubstituted products, depending on reaction conditions.

L4 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN

AN 1968:435972 CAPLUS

DN 69:35972

TI 4-Methylisoquinolines

IN Aebi, Albert; Nair, Mohan D.; Bucher, Karl

PA CIBA Ltd.

SO Patentschrift (Switz.), 6 pp.

CODEN: SWXXAS

DT Patent

LA German

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	CH 438308		19671215	CH	19630221

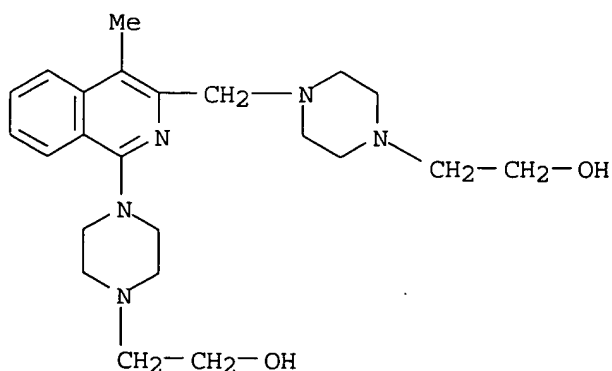
IT **14576-16-0P 14576-17-1P 14577-67-4P**

14825-52-6P 18704-43-3P

RL: SPN (Synthetic preparation); PREP (Preparation)
(prepn. of)

RN 14576-16-0 CAPLUS

CN 1-Piperazineethanol, 4-[[1-[4-(2-hydroxyethyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]- (9CI) (CA INDEX NAME)



RN 14576-17-1 CAPLUS

CN 1-Piperazinecarboxylic acid, 4-[[1-[4-(ethoxycarbonyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]-, ethyl ester (9CI) (CA INDEX NAME)

L4 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN

AN 1968:435972 CAPLUS

DN 69:35972

TI 4-Methylisoquinolines

IN Aebl, Albert; Nair, Mohan D.; Bucher, Karl

PA CIBA Ltd.

SO Patentschrift (Switz.), 6 pp.

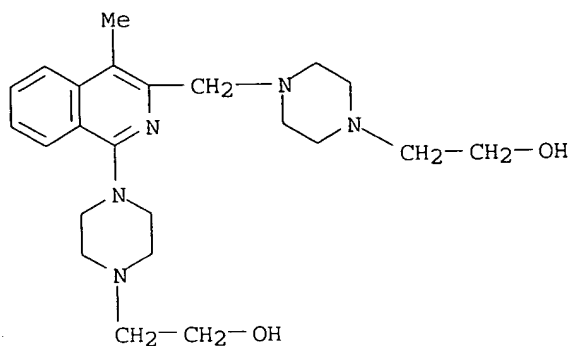
CODEN: SWXXAS

DT Patent

LA German

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	CH 438308		19671215	CH	19630221
IT	14576-16-0P 14576-17-1P 14577-67-4P 14825-52-6P 18704-43-3P				
	RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of)				
RN	14576-16-0	CAPLUS			
CN	1-Piperazineethanol, 4-[[1-[4-(2-hydroxyethyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]- (9CI) (CA INDEX NAME)				

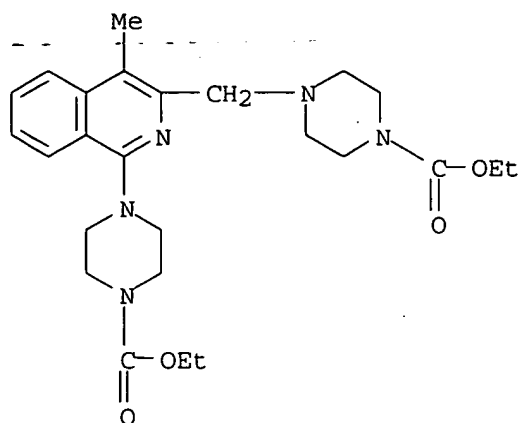


RN 14576-17-1 CAPLUS

CN 1-Piperazinecarboxylic acid, 4-[[1-[4-(ethoxycarbonyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]-, ethyl ester (9CI) (CA INDEX NAME)

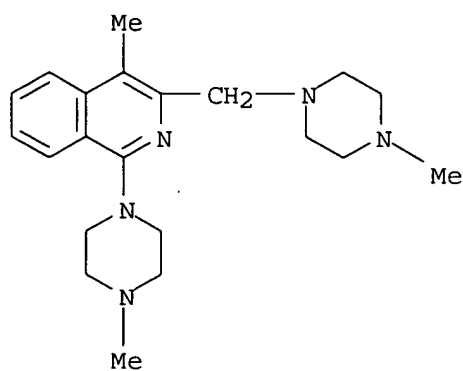
Patel

<7/28/2003>



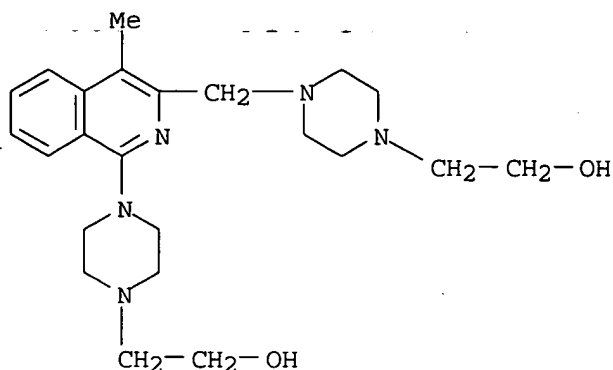
RN 14577-67-4 CAPLUS

CN Isoquinoline, 4-methyl-1-(4-methyl-1-piperazinyl)-3-[(4-methyl-1-piperazinyl)methyl]- (8CI, 9CI) (CA INDEX NAME)



RN 14825-52-6 CAPLUS

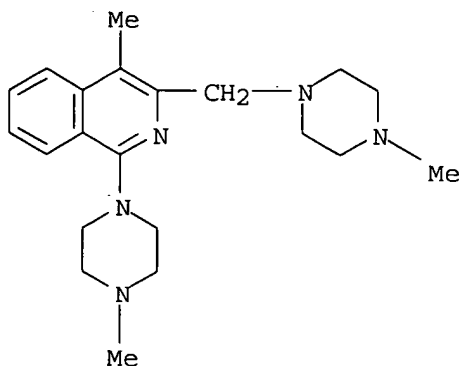
CN 1-Piperazineethanol, 4,4'-[methylene(4-methyl-3,1-isoquinolinediyl)]di-, hydrochloride (8CI) (CA INDEX NAME)



● x HCl

RN 18704-43-3 CAPLUS

CN Isoquinoline, 4-methyl-1-((4-methyl-1-piperazinyl)methyl)-3-((4-methyl-1-piperazinyl)methyl)-, monohydrochloride (8CI) (CA INDEX NAME)



● HCl

GI For diagram(s), see printed CA Issue.

AB The title compds. are prepd. by treating 1-chloro-3-chloromethyl-4-methylisoquinoline (I) or its substituted derivs. with secondary amines. Thus, 1.55 g. I and 5 ml. morpholine was heated overnight in a pressure vessel at 150.degree.. The cryst. suspension was then evapd. to dryness, taken up in CHCl₃, extd. 2 times with dil. aq. HCl, and the aq. exts. adjusted to pH 8-9 with NaOH. The oil which sepd. gradually crystd., and was sepd. and recrystd. from iso-PROH to give II (R = H and R₁ = morpholino), m. 100.degree.; dihydrochloride m. 229-32.degree. (decompn.) and maleate m. 173-5.degree.. Other II similarly prepd. are shown in the table. The starting material for II (R = NO₂) was prepd. by treating I with concd. H₂SO₄ and fuming HNO₃ to give II (R = NO₂, R₁ = Cl), m. 104-5.degree.. A mixt. of 4 g. 1,7-dichloro-3-chloromethyl-4-methylisoquinoline (IV) and 50 ml. morpholine was refluxed 4 hrs., and excess morpholine was then removed under reduced pressure. [TABLE

OMITTED] The residue was treated with aq. Na₂CO₃ until alk. and extd. with CHCl₃. The exts. were evapd. to give 7-chloro-4-methyl-1-morpholino-3-(morpholinomethyl)isoquinoline, which was purified by conversion to its maleate and then to the free base, m. 120.degree. (EtOH). IV was prepd. by treating 4,4-dimethylhomophthalimide with fuming HNO₃ and concd. H₂SO₄ at -10.degree. to give 4,4-dimethyl-7-nitrohomophthalimide, m. 209-11.degree.. Hydrogenation over Pd-C gave the 7-amino compd., m. 176-9.degree., which was diazotized and treated with CuCl to give the 7-chloro deriv., m. 200.degree.. Treatment with POCl₃ gave IV, m. 135.degree.. These compds. are used in pharmaceutical applications.

L4 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN

AN 1967:421848 CAPLUS

DN 67:21848

TI New antitussive isoquinoline derivatives

PA CIBA Ltd.

SO Fr. M., 10 pp.

CODEN: FMXXAJ

DT Patent

LA French

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	FR 3782		19660131		
				CH	19630121
				CH	19640121

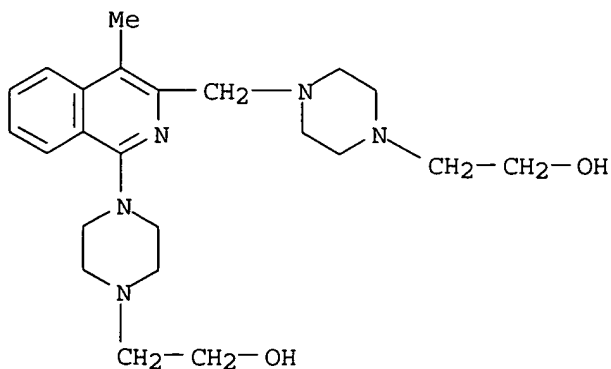
IT 14576-16-0P 14576-17-1P 14577-67-4P

14601-04-8P 14825-52-6P

RL: SPN (Synthetic preparation); PREP (Preparation)
(prepn. of)

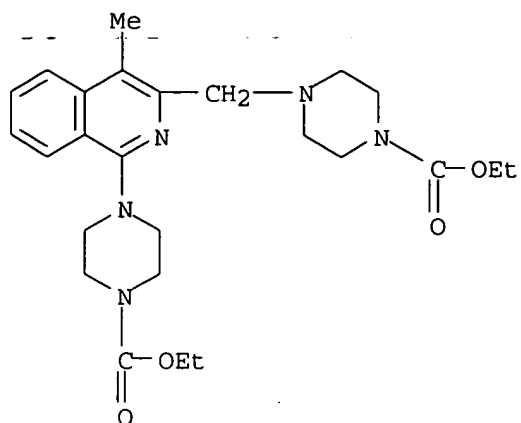
RN 14576-16-0 CAPLUS

CN 1-Piperazineethanol, 4-[[1-[4-(2-hydroxyethyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]- (9CI) (CA INDEX NAME)



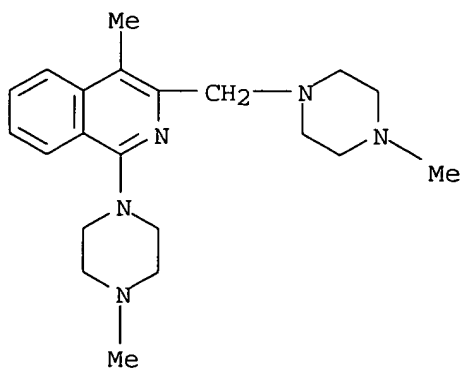
RN 14576-17-1 CAPLUS

CN 1-Piperazinecarboxylic acid, 4-[[1-[4-(ethoxycarbonyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]-, ethyl ester (9CI) (CA INDEX NAME)



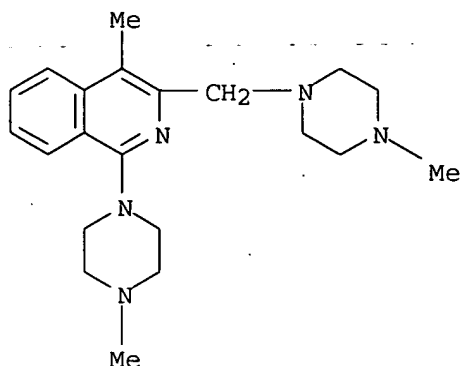
RN 14577-67-4 CAPLUS

CN Isoquinoline, 4-methyl-1-((4-methyl-1-piperazinyl)methyl)-3-((4-methyl-1-piperazinyl)methyl)carbamate (8CI, 9CI) (CA INDEX NAME)



RN 14601-04-8 CAPLUS

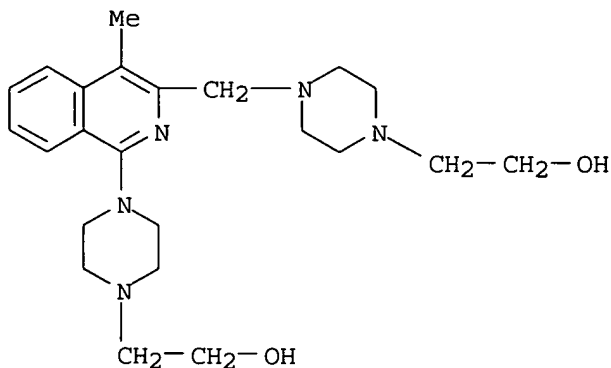
CN Isoquinoline, 4-methyl-1-((4-methyl-1-piperazinyl)methyl)-3-((4-methyl-1-piperazinyl)methyl)carbamate (8CI) (CA INDEX NAME)



●x HCl

RN 14825-52-6 CAPLUS

CN 1-Piperazineethanol, 4,4'-[methylene(4-methyl-3,1-isoquinolinediyl)]di-, hydrochloride (8CI) (CA INDEX NAME)



●x HCl

GI For diagram(s), see printed CA Issue.

AB New antitussive isoquinoline derivs. with general formula (I) are prepd. A mixt. of 9 g. 1-chloro-3-chloromethyl-4-methylisoquinoline (II) and 40 cc. piperidine (III) is heated in a sealed tube 8 hrs. at 150.degree., the reaction mixt. concd. in vacuo, treated with water, and extd. with CH₂Cl₂, the ext. dried and evapd. to dryness, and the residue in CHCl₃ passed through activated alumina to give 4-methyl-1-piperidino-3-piperidinomethylisoquinoline, m. 111.degree. (water-EtOH). The following products are prepd. in a similar way (starting materials, reaction time, reaction temp., final product, m.p., derivs., and m.p. given): II (9 g.), pyrrolidine (40 cc.), 8 hrs., 150.degree., 4-methyl-1-(1-pyrrolidinyl)-3-(1-pyrrolidinylmethyl)isoquinoline, -, hydrochloride, 239.degree.; II (8 g.), N-methylpiperazine (IV) (50 cc.), 8 hrs., 150.degree., 4-methyl-1-(N'-methylpiperazino)-3-(N'-methylpiperazinomethyl)isoquinoline, 110-11.degree., hydrochloride, 238.degree.; II (8 g.),

N-(.beta.-hydroxyethyl)piperazine (40 cc.), 8 hrs., 150.degree., 4-methyl-1-[N'-(.beta.-hydroxyethyl)piperazino]-3-[N'-(.beta.-hydroxyethyl)piperazinomethyl]isoquinoline, 112.degree., hydrochloride, 262.degree. (decompn.); II (6 g.), Et₂NH (15 cc.), 8 hrs., 150.degree., 4-methyl-1-diethylamino-3-diethylaminomethylisoquinoline, -, dimaleate, 109-11.degree.; II (4.5 g.), ethanolamine (15 cc.), 3 hrs., 130.degree., 4-methyl-1-(.beta.-hydroxyethylamino)-3-(.beta.-hydroxyethylaminomethyl)isoquinoline, -, hydrochloride, 252-4.degree.; II (5 g.), N-carbethoxypiperazine (V) (20 cc.), 6 hrs., 140.degree., 4-methyl-1-(N'-carbethoxypiperazino)-3-(N'-carbethoxypiperazinomethyl)isoquinoline, 90-2.degree., -, -; II (5 g.), 2-methylpiperidine (20 cc.), 6 hrs., 140.degree., 1-chloro-4-methyl-3-(2-methylpiperidinomethyl)isoquinoline (VI), 106-8.degree., -, -; VI (6 g.), morpholine (VII) (20 cc.), 14 hrs., 170.degree., 4-methyl-1-morpholino-3-(2-methylpiperidinomethyl)isoquinoline, 103-4.degree., -, -; 1-chloro-3-chloromethyl-4-methyl-5-nitroisoquinoline (VIII) (2 g.), VII (10 cc.), 2 hrs., 120.degree., 4-methyl-1-morpholino-3-morpholinomethyl-5-nitroisoquinoline (IX), 145-6.degree., -, -; VIII (2.5 g.), III (10 cc.), 2.5 hrs., 80.degree., 4-methyl-5-nitro-1-piperidino-3-piperidinomethylisoquinoline, 104-6.degree., -, -; VIII (2.5 g.), p-anisidine (4.55 g.), EtOH (80 cc.), 4 hrs., reflux, 1-p-anisidino-3-p-anisidinomethyl-4-methyl-5-nitroisoquinoline, 183-5.degree., -, -; 1,7-dichloro-3-chloromethyl-4-methylisoquinoline (X) (4 g.), VII (50 cc.), 4 hrs., reflux, 7-chloro-4-methyl-1-morpholino-3-morpholinomethylisoquinoline, 120.degree., maleate, -, -; VIII (5 g.), III (8 cc.), EtOH (75 cc.), 1 hr., reflux, 1-chloro-4-methyl-5-nitro-3-piperidinomethylisoquinoline, 67-79.degree., -, -; II (4.5 g.), III (15 cc.), 2 hrs., 80.degree., 1-chloro-4-methyl-3-piperidinomethylisoquinoline, 79-80.degree., -, -; VIII (3.5 g.), IV (2.58 g.), EtOH (100 cc.), 2 hrs., reflux, 1-chloro-3-(N'-methylpiperazinomethyl)-4-methyl-5-nitroisoquinoline, 173-5.degree., -, -; VIII (4 g.), V (10 cc.), EtOH (75 cc.), 1 hr., reflux, 1-chloro-3-(N'-carbethoxypiperazinomethyl)-4-methyl-5-nitroisoquinoline, 127-8.degree., -, -; VIII (2.71 g.), diethanolamine (4.5 g.), dioxane (50 cc.), 3 hrs., reflux, 1-chloro-3-[bis(.beta.-hydroxyethyl)aminomethyl]-4-methyl-5-nitroisoquinoline, 110-12.degree., -, -; II (5.0 g.), 4-methylpiperidine (5.5 cc.), 2 hrs., 80.degree., 1-chloro-3-(4-methylpiperidinomethyl)-4-methylisoquinoline, 83-5.degree., -, -; II (5.0 g.), concd. aq. NH₃ (80 cc.), hydrated CuSO₄ (1.0 g.), 30 hrs., 140.degree., bis(1-chloro-4-methyl-3-isoquinolylmethyl)amine, 131-2.degree., -, -; II (5.0 g.), N-(.gamma.-aminopropyl)morpholine (6.5 g.), 2 hrs., 100.degree., N,N-bis(1-chloro-4-methyl-3-isoquinolylmethyl)-N-(.gamma.-morpholinopropyl)amine, 110-12.degree., -, -. Some starting materials and other products are prepd. as follows: II (6 g.) is added slowly with stirring to a cooled mixt. of 15 cc. concd. H₂SO₄ and 15 cc. fuming HNO₃ and the mixt. stirred 1.5 hrs. below 5.degree. and poured over a mixt. of ice and water to ppt. VIII, m 104-5.degree. (EtOH). A mixt. of 4 g. IX, 0.3 g. Pd-C and 150 cc. 95% EtOH is hydrogenated 1.5 hrs. to give 5-amino-4-methyl-1-morpholino-3-morpholinomethylisoquinoline (XI), m. 134-5.degree. (EtOH). A soln. of 1.6 g. NaNO₂ in 5 cc. water is added slowly to a cooled soln. of 8 g. XI in 6 cc. concd. HCl and 6 cc. water, the resulting soln. poured into a cooled soln. of Cu₂Cl₂ (prepd. from 8 g. CuSO₄) and then is heated at 60.degree., and the ppt. suspended in 25 cc. water, alkalized, and extd. with CHCl₃ to give 5-chloro-4-methyl-1-morpholino-3-morpholinomethylisoquinoline, m. 104.degree.. 4,4-Dimethylhomophthalimide (15 g.) is added slowly with stirring to a cooled (-10.degree.) mixt. of 30 cc. concd. H₂SO₄ and 30 cc. fuming HNO₃ and the mixt. stirred 1 hr. below 20.degree. and poured over a mixt. of ice and

water to ppt. 4,4-dimethyl-7-nitrohomophthalimide (XII), m. 209-11.degree. (EtOH). A mixt. of 23.4 g. XII, 0.5 g. Pd-C, and 200 cc. MeOH is hydrogenated at 50.degree./3.4 atm. .apprx.1.5 hrs. to give 4,4-dimethyl-7-aminohomophthalimide (XIII), m. 176-9.degree. (MeOH). Concd. H₂SO₄ (26 g.) is added slowly to a mixt. of 20 g. XIII and 90 cc. water, and cooled at 0.degree., 8.4 g. NaNO₂ in 24 cc. water added slowly to it, and this mixt. is added slowly to a soln. of Cu₂Cl₂ (prepd. from 33.4 g. CuSO₄), and the mixt. heated at 60.degree. 30 min., cooled, dild. with water, and extd. with CHCl₃ to give 4,4-dimethyl-7-chlorohomophthalimide (XIV), m. 200.degree. (EtOH). A mixt. of 10 g. XIV, 0.5 cc. water, and 40 cc. POCl₃ is heated in a sealed tube at 200.degree. 5 hrs. to give X, m. 135.degree. (hexane-CHCl₃). Some recipes for the prepn. of pharmacol. compns. are also given.

L Number	Hits	Search Text	DB	Time stamp
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